FILE 'HOME' ENTERED AT 15:40:40 ON 24 MAR 2006

=> file .mymstn

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

0.21

FULL ESTIMATED COST 0.21

FILE 'MEDLINE' ENTERED AT 15:40:53 ON 24 MAR 2006

FILE 'AGRICOLA' ENTERED AT 15:40:53 ON 24 MAR 2006

FILE 'JICST-EPLUS' ENTERED AT 15:40:53 ON 24 MAR 2006

COPYRIGHT (C) 2006 Japan Science and Technology Agency (JST)

FILE 'BIOSIS' ENTERED AT 15:40:53 ON 24 MAR 2006

Copyright (c) 2006 The Thomson Corporation

FILE 'CAPLUS' ENTERED AT 15:40:53 ON 24 MAR 2006

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'LIFESCI' ENTERED AT 15:40:53 ON 24 MAR 2006

COPYRIGHT (C) 2006 Cambridge Scientific Abstracts (CSA)

FILE 'BIOTECHNO' ENTERED AT 15:40:53 ON 24 MAR 2006 COPYRIGHT (C) 2006 Elsevier Science B.V., Amsterdam. All rights reserved.

FILE 'EMBASE' ENTERED AT 15:40:53 ON 24 MAR 2006

Copyright (c) 2006 Elsevier B.V. All rights reserved.

=> s adzyme

L1 2 ADZYME

=> s fusion (w) protein and catalytic (w) domain and target and targeting (w) domain L2 11 FUSION (W) PROTEIN AND CATALYTIC (W) DOMAIN AND TARGET AND TARGE TING (W) DOMAIN

 $\Rightarrow$  s chimeric (w) protein and catalytic (w) domain and target and targeting (w) domain

L3 0 CHIMERIC (W) PROTEIN AND CATALYTIC (W) DOMAIN AND TARGET AND TARGETING (W) DOMAIN

=> d ibib abs 12 1-11

L2 ANSWER 1 OF 11 MEDLINE on STN ACCESSION NUMBER: 95318049 MEDLINE DOCUMENT NUMBER: PubMed ID: 7541032

DOCUMENT NUMBER: PubMe TITLE: T cel

T cell-targeted immunofusion proteins from Escherichia

coli.

AUTHOR: Better M; Bernhard S L; Williams R E; Leigh S D; Bauer R J;

Kung A H; Carroll S F; Fishwild D M

CORPORATE SOURCE: XOM

XOMA Corporation, Santa Monica, California 90404, USA. The Journal of biological chemistry, (1995 Jun 23) Vol.

270, No. 25, pp. 14951-7.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

SOURCE:

United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199507

ENTRY DATE: Entered STN: 19950817

## Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID: ssptamym1652

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

```
Web Page URLs for STN Seminar Schedule - N. America
NEWS
     1
NEWS
     2
                 "Ask CAS" for self-help around the clock
                IPC search and display fields enhanced in CA/CAplus with the
        DEC 21
NEWS
                 IPC reform
                New IPC8 SEARCH, DISPLAY, and SELECT fields in USPATFULL/
NEWS
        DEC 23
     4
                 USPAT2
NEWS 5
        JAN 13
                IPC 8 searching in IFIPAT, IFIUDB, and IFICDB
NEWS 6
                New IPC 8 SEARCH, DISPLAY, and SELECT enhancements added to
        JAN 13
                 INPADOC
NEWS 7
        JAN 17
                Pre-1988 INPI data added to MARPAT
NEWS 8
                IPC 8 in the WPI family of databases including WPIFV
        JAN 17
NEWS 9 JAN 30
                Saved answer limit increased
                Monthly current-awareness alert (SDI) frequency
NEWS 10 JAN 31
                 added to TULSA
                STN AnaVist, Version 1.1, lets you share your STN AnaVist
NEWS 11 FEB 21
                 visualization results
NEWS 12 FEB 22
                 Status of current WO (PCT) information on STN
                The IPC thesaurus added to additional patent databases on STN
NEWS 13 FEB 22
                Updates in EPFULL; IPC 8 enhancements added
NEWS 14 FEB 22
                New STN AnaVist pricing effective March 1, 2006
NEWS 15 FEB 27
                MEDLINE/LMEDLINE reload improves functionality
NEWS 16 FEB 28
        FEB 28
                TOXCENTER reloaded with enhancements
NEWS 17
NEWS 18 FEB 28
                REGISTRY/ZREGISTRY enhanced with more experimental spectral
                 property data
NEWS 19 MAR 01
                 INSPEC reloaded and enhanced
                Updates in PATDPA; addition of IPC 8 data without attributes
        MAR 03
NEWS 20
        MAR 08
                X.25 communication option no longer available after June 2006
NEWS 21
NEWS 22 MAR 22
                EMBASE is now updated on a daily basis
```

NEWS EXPRESS FEBRUARY 15 CURRENT VERSION FOR WINDOWS IS V8.01a,
CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
AND CURRENT DISCOVER FILE IS DATED 19 DECEMBER 2005.
V8.0 AND V8.01 USERS CAN OBTAIN THE UPGRADE TO V8.01a AT
http://download.cas.org/express/v8.0-Discover/

NEWS HOURS STN Operating Hours Plus Help Desk Availability NEWS LOGIN Welcome Banner and News Items

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

Last Updated on STN: 19960129 Entered Medline: 19950731

Fusion proteins between cell-targeting AB domains and cytotoxic proteins should be particularly effective therapeutic reagents. We constructed a family of immunofusion proteins linking humanized Fab, F(ab')2, or single chain antibody forms of the H65 antibody (which recognizes the CD5 antigen on the surface of human T cells) with the plant ribosome-inactivating protein gelonin. We reasoned that such an immunofusion would kill human target cells as efficiently as the previously described chemical conjugates of H65 and gelonin (Better M., Bernhard, S. L., Fishwild, D. M., Nolan, P. A., Bauer, R. J., Kung, A. H. C., and Carroll, S. F. (1994) J. Biol. Chemical 269, 9644-9650) if both the recognition and catalytic domains remained active, and a proper linkage between domains could be found. Immunofusion proteins were produced in Escherichia coli as secreted proteins and were recovered directly from the bacterial culture supernatant in an active form. All of the immunofusion proteins were purified by a common process and were tested for cytotoxicity toward antigen-positive human cells. A 20-60-fold range of cytotoxic activity was seen among the fusion family members, and several fusion proteins were identified which are approximately as active as effective chemical conjugates. Based on these constructs, immunofusion avidity and potency can be controlled by appropriate selection of antibody domains and ribosome-inactivating protein.

L2 ANSWER 2 OF 11 MEDLINE ON STN ACCESSION NUMBER: 94224796 MEDLINE DOCUMENT NUMBER: PubMed ID: 8170960

TITLE: Functionally active targeting domain of

the beta-adrenergic receptor kinase: an inhibitor of G beta

gamma-mediated stimulation of type II adenylyl cyclase.

AUTHOR: Inglese J; Luttrell L M; Iniguez-Lluhi J A; Touhara K; Koch

W J; Lefkowitz R J

CORPORATE SOURCE: Department of Medicine, Howard Hughes Medical Institute,

Duke University Medical Center, Durham, NC 27710.

CONTRACT NUMBER: HL16037 (NHLBI)

SOURCE: Proceedings of the National Academy of Sciences of the

United States of America, (1994 Apr 26) Vol. 91, No. 9, pp.

3637-41.

Journal code: 7505876. ISSN: 0027-8424.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199406

ENTRY DATE: Entered STN: 19940613

Last Updated on STN: 20021218 Entered Medline: 19940601

The beta-adrenergic receptor kinase (beta ARK) phosphorylates its AB membrane-associated receptor substrates, such as the beta-adrenergic receptor, triggering events leading to receptor desensitization. beta ARK activity is markedly stimulated by the isoprenylated beta gamma subunit complex of heterotrimeric guanine nucleotide-binding proteins (G beta gamma), which translocates the kinase to the plasma membrane and thereby targets it to its receptor substrate. The amino-terminal two-thirds of beta ARK1 composes the receptor recognition and catalytic domains, while the carboxyl third contains the G beta gamma binding sequences, the targeting domain. We prepared this domain as a recombinant His6 fusion protein from Escherichia coli and found that it had both independent secondary structure and functional activity. We demonstrated the inhibitory properties of this domain against G beta gamma activation of type II adenylyl cyclase both in a reconstituted system utilizing Sf9 insect cell membranes and in a permeabilized 293 human embryonic kidney

cell system. Gi alpha-mediated inhibition of adenylyl cyclase was not affected. These data suggest that this His6 fusion protein derived from the carboxyl terminus of beta ARK1 provides a specific probe for defining G beta gamma-mediated processes and for studying the structural features of a G beta gamma-binding domain.

22 ANSWER 3 OF 11 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN

ACCESSION NUMBER: 1995:362428 BIOSIS DOCUMENT NUMBER: PREV199598376728

TITLE: T Cell-targeted Immunofusion Proteins from Escherichia

coli.

AUTHOR(S): Better, Marc [Reprint author]; Bernhard, Susan L.;

Williams, Robert E.; Leigh, Scott D.; Bauer, Robert J.; Kung, Ada H. C.; Carroll, Stephen F.; Fishwild, Dianne M. Xoma Corp., 1545 17th St., Santa Monica, CA 90404, USA

SOURCE: Journal of Biological Chemistry, (1995) Vol. 270, No. 25,

pp. 14951-14957.

CODEN: JB

CODEN: JBCHA3. ISSN: 0021-9258.

DOCUMENT TYPE: Article LANGUAGE: English

CORPORATE SOURCE:

ENTRY DATE: Entered STN: 30 Aug 1995

Last Updated on STN: 30 Aug 1995

AB Fusion proteins between cell-targeting

domains and cytotoxic proteins should be particularly effective therapeutic reagents. We constructed a family of immunofusion proteins linking humanized Fab, F(ab')-2, or single chain antibody forms of the H65 antibody (which recognizes the CD5 antigen on the surface of human T cells) with the plant ribosome-inactivating protein gelonin. We reasoned that such an immunofusion would kill human target cells as efficiently as the previously described chemical conjugates of H65 and gelonin (Better M., Bernhard, S. L., Fishwild, D. M., Nolan, P. A., Bauer, R. J., Kung, A. H. C., and Carroll, S. F. (1994) J. Biol. Chemical 269, 9644-9650) if both the recognition and catalytic domains remained active, and a proper linkage between domains could be found. Immunofusion proteins were produced in Escherichia coli as secreted proteins and were recovered directly from the bacterial culture supernatant in an active form. All of the immunofusion proteins were purified by a common process and were tested for cytotoxicity toward antigen-positive human cells. A 20-60-fold range of cytotoxic activity was seen among the fusion family members, and several fusion proteins were identified which are approximately as active as effective chemical conjugates. Based on these constructs, immunofusion avidity and potency can be controlled by appropriate selection of antibody domains and ribosome-inactivating protein.

L2 ANSWER 4 OF 11 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN

ACCESSION NUMBER: 1994:304464 BIOSIS DOCUMENT NUMBER: PREV199497317464

TITLE: Functionally active targeting domain of

the beta-adrenergic receptor kinase: An inhibitor of G-beta-gamma-mediated stimulation of type II adenylyl

cyclase.

AUTHOR(S): Inglese, J. [Reprint author]; Luttrell, L. M. [Reprint

author]; Iniguez-Lluhi, J. A.; Touhara, K. [Reprint author]; Koch, W. J. [Reprint author]; Lefkowitz, R. J.

[Reprint author]

CORPORATE SOURCE: Dep. Med., Box 3821, Howard Hughes Med. Inst., Duke

University Med. Center, Durham, NC 27710, USA

SOURCE: Proceedings of the National Academy of Sciences of the

United States of America, (1994) Vol. 91, No. 9, pp.

3637-3641.

CODEN: PNASA6. ISSN: 0027-8424.

DOCUMENT TYPE: Article LANGUAGE: English

ENTRY DATE: Entered STN: 13 Jul 1994

Last Updated on STN: 24 Aug 1994

The beta-adrenergic receptor kinase (beta-ARK) phosphorylates its AB membrane-associated receptor substrates, such as the beta-adrenergic receptor, triggering events leading to receptor desensitization. beta-ARK activity is markedly stimulated by the isoprenylated beta-gamma subunit complex of heterotrimeric guanine nucleotide-binding proteins (G-beta-gamma), which translocates the kinase to the plasma membrane and thereby targets it to its receptor substrate. The amino-terminal two-thirds of beta-ARK1 composes the receptor recognition and catalytic domains, while the carboxyl third contains the G-beta-gamma binding sequences, the targeting domain. We prepared this domain as a recombinant His-6 fusion protein from Escherichia coli and found that it had both independent secondary structure and functional activity. We demonstrated the inhibitory properties of this domain against G-beta-gamma activation of type II adenylyl cyclase both in a reconstituted system utilizing Sf9 insect cell membranes and in a permeabilized 293 human embryonic kidney cell system. G-ialpha-mediated inhibition of adenylyl cyclase was not affected. These data suggest that this His-6 fusion protein derived from the carboxyl terminus of beta-ARK1 provides a specific probe for defining G-beta-gamma-mediated processes and for studying the structural features of a G-beta-gamma-binding domain.

L2 ANSWER 5 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1995:656920 CAPLUS

DOCUMENT NUMBER: 123:81252

TITLE: T cell-targeted immunofusion proteins from Escherichia

coli

AUTHOR(S): Better, Marc; Bernhard, Susan L.; Williams, Robert E.;

Leigh, Scott D.; Bauer, Robert J.; Kung, Ada H. C.;

Carroll, Stephen F.; Fishwild, Dianne M.

CORPORATE SOURCE: XOMA Corp., Santa Monica, CA, 90404, USA

SOURCE: Journal of Biological Chemistry (1995), 270(25),

14951-7

CODEN: JBCHA3; ISSN: 0021-9258

PUBLISHER: American Society for Biochemistry and Molecular Bio

logy

DOCUMENT TYPE: Journal LANGUAGE: English

AB Fusion proteins between cell-targeting

domains and cytotoxic proteins should be particularly effective therapeutic reagents. We constructed a family of immunofusion proteins linking humanized Fab, F(ab')2, or single chain antibody forms of the H65 antibody (which recognizes the CD5 antigen on the surface of human T cells) with the plant ribosome-inactivating protein gelonin. We reasoned that such an immunofusion would kill human target cells as efficiently as the previously described chemical conjugates of H65 and gelonin if both the recognition and catalytic domains remained active, and a proper linkage between domains could be found. Immunofusion proteins were produced in Escherichia coli as secreted proteins and were recovered directly from the bacterial culture supernatant in an active form. All of the immunofusion proteins were purified by a common process and were tested for cytotoxicity toward antigen-pos. human cells. A 20-60-fold range of cytotoxic activity was seen among the fusion family members, and several fusion proteins were identified which are approx. as active as effective chemical conjugates. Based on these constructs, immunofusion avidity and potency can be controlled by appropriate selection of antibody domains and ribosome-inactivating protein.

L2 ANSWER 6 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN ACCESSION NUMBER: 1994:502907 CAPLUS

DOCUMENT NUMBER: 121:102907

TITLE: Functionally active targeting domain

of the  $\beta$ -adrenergic receptor kinase: an inhibitor

of GBy-mediated stimulation of type II

adenylyl cyclase

AUTHOR(S): Inglese, J.; Luttrell, L. M.; Iniguez-Lluhi, J. A.;

Touhara, K.; Koch, W. J.; Lefkowitz, R. J.

CORPORATE SOURCE: Howard Hughes Med. Inst., Duke Univ., Durham, NC,

27710, USA

SOURCE: Proceedings of the National Academy of Sciences of the

United States of America (1994), 91(9), 3637-41

CODEN: PNASA6; ISSN: 0027-8424

DOCUMENT TYPE: Journal LANGUAGE: English

The  $\beta$ -adrenergic receptor kinase ( $\beta$ ARK) phosphorylates its membrane-associated receptor substrates, such as the  $\beta$ -adrenergic receptor, triggering events leading to receptor desensitization. The βARK activity is markedly stimulated by the isopropenylated βγ subunit complex of heterotrimeric guanine nucleotide-binding proteins (Gβγ), which translocates the kinase to the plasma membrane and thereby targets it to its receptor subunits. The amino-terminal two-thirds of  $\beta$ ARK1 composes the receptor recognition and catalytic domains, while the carboxyl third contains the  $G\beta\gamma$  binding sequences, the targeting domain. The authors prepared this domain as a recombinant His6 fusion protein from Escherichia coli and found that it had both independent second structure and functional activity. The authors demonstrated the inhibitory properties of this domain against Gβγ activation of type II adenylyl cyclase both in a reconstituted system utilizing Sf9 insect cell membranes and in a permeabilized 293 human embryonic kidney cell system.  $Gl\alpha$ -mediated inhibition of adenylyl cyclase was not affected. These data suggest that this His6 fusion protein derived from the carboxyl terminus of BARK1 provides a specific probe for defining  $G\beta\gamma$ -mediated processes and for studying the structural features of a  $G\beta\gamma$ -binding domain.

L2 ANSWER 7 OF 11 LIFESCI COPYRIGHT 2006 CSA on STN

ACCESSION NUMBER: 96:19231 LIFESCI

TITLE: T cell-targeted immunofusion proteins from Escherichia coli
AUTHOR: Better, M.; Bernhard, S.L.; Williams, R.E.; Leigh, S.D.;
Bauer, R.J.; Kung, A.H.C.; Carroll, S.F.; Fishwild, D.M.

CORPORATE SOURCE: Xoma Corp., 1545 17th St., Santa Monica, CA 90404, USA SOURCE: J. BIOL. CHEM., (1995) vol. 270, no. 25, pp. 14591-14597.

ISSN: 0021-9528.

DOCUMENT TYPE: Journal FILE SEGMENT: F; W3 LANGUAGE: English SUMMARY LANGUAGE: English

AB Fusion proteins between cell-targeting

domains and cytotoxic proteins should be particularly effective therapeutic reagents. We constructed a family of immunofusion proteins linking humanized Fab, F(ab') sub(2), or single chain antibody forms of the H65 antibody (which recognizes the CD5 antigen on the surface of human T cells) with the plant ribosome-inactivating protein gelonin. We reasoned that such an immunofusion would kill human target cells as efficiently as the previously described chemical conjugates of H65 and gelonin if both the recognition and catalytic domains remained active, and a proper linkage between domains could be found. Immunofusion proteins were produced in Escherichia coli as secreted proteins and were recovered directly from the bacterial culture supernatant in an active form. All of the immunofusion proteins were purified by a common process and were tested for cytotoxicity toward antigen-positive human cells. A 20-60-fold range of cytotoxic activity was

seen among the fusion family members, and several fusion proteins were identified which are ap proximately as active as effective chemical conjugates. Based on these constructs, immunofusion avidity and potency can be controlled by appropriate selection of antibody domains and ribosome-inactivating protein.

L2 ANSWER 8 OF 11 BIOTECHNO COPYRIGHT 2006 Elsevier Science B.V. on STN

ACCESSION NUMBER: 1995:25189002 BIOTECHNO

TITLE: T cell-targeted immunofusion proteins from Escherichia

coli

AUTHOR: Better M.; Bernhard S.L.; Williams R.E.; Leigh S.D.;

Bauer R.J.; Kung A.H.C.; Carroll S.F.; Fishwild D.M.

CORPORATE SOURCE: Xoma Corp., 1545 17th St., Santa Monica, CA 90404,

United States.

SOURCE: Journal of Biological Chemistry, (1995), 270/25

(14951-14957)

CODEN: JBCHA3 ISSN: 0021-9258

DOCUMENT TYPE: Journal; Article COUNTRY: United States

LANGUAGE: English
SUMMARY LANGUAGE: English
AN 1995:25189002 BIOTECHNO

AB Fusion proteins between cell-targeting

domains and cytotoxic proteins should be particularly effective therapeutic reagents. We constructed a family of immunofusion proteins linking humanized Fab, F(ab').sub.2, or single chain antibody forms of the H65 antibody (which recognizes the CD5 antigen on the surface of human T cells) with the plant ribosome-inactivating protein gelonin. We reasoned that such an immunofusion would kill human target cells as efficiently as the previously described chemical conjugates of H65 and gelonin (Better M., Bernhard, S. L., Fishwild, D. M., Nolan, P. A., Bauer, R. J., Kung, A. H. C., and Carroll, S. F. (1994) J. Biol. Chemical 269, 9644-9650) if both the recognition and catalytic domains remained active, and a proper linkage between domains could be found. Immunofusion proteins were produced in Escherichia coli as secreted proteins and were recovered directly from the bacterial culture supernatant in an active form. All of the immunofusion proteins were purified by a common process and were tested for cytotoxicity toward antigen-positive human cells. A 20-60-fold range of cytotoxic activity was seen among the fusion family members, and several fusion proteins were identified which are approximately as active as effective chemical conjugates. Based on these constructs, immunofusion avidity and potency can be controlled by appropriate selection of antibody domains and ribosome- inactivating protein.

L2 ANSWER 9 OF 11 BIOTECHNO COPYRIGHT 2006 Elsevier Science B.V. on STN

ACCESSION NUMBER: 1994:24139486 BIOTECHNO

TITLE: Functionally active targeting domain

of the  $\beta$ -adrenergic receptor kinase: An inhibitor

of  $G(\beta\gamma)$ -mediated stimulation of type II

adenylyl cyclase

AUTHOR: Inglese J.; Luttrell L.M.; Iniguez-Lluhi J.A.; Touhara

K.; Koch W.J.; Lefkowitz R.J.

CORPORATE SOURCE: Department of Medicine, Howard Hughes Medical

Institute, Duke University Medical Center, Durham, NC

27710, United States.

SOURCE: Proceedings of the National Academy of Sciences of the

United States of America, (1994), 91/9 (3637-3641)

CODEN: PNASA6 ISSN: 0027-8424

DOCUMENT TYPE: Journal; Article

COUNTRY: United States LANGUAGE: English

SUMMARY LANGUAGE: English AN 1994:24139486 BIOTECHNO

The  $\beta$ -adrenergic receptor kinase ( $\beta$ ARK) phosphorylates its AB membrane- associated receptor substrates, such as the  $\beta$ -adrenergic receptor, triggering events leading to receptor desensitization. βARK activity is markedly stimulated by the isoprenylated βy subunit complex of heterotrimeric guanine nucleotide-binding proteins  $(G(\beta\gamma))$ , which translocates the kinase to the plasma membrane and thereby targets it to its receptor substrate. The amino-terminal two-thirds of βARK1 composes the receptor recognition and catalytic domains, while the carboxyl third contains the  $G(\beta y)$  binding sequences, the targeting domain. We prepared this domain as a recombinant His.sub.6 fusion protein from Escherichia coli and found that it had both independent secondary structure and functional activity. We demonstrated the inhibitory properties of this domain against  $G(\beta \gamma)$  activation of type II adenylyl cyclase both in a reconstituted system utilizing Sf9 insect cell membranes and in a permeabilized 293 human embryonic kidney cell system.  $G(i\alpha)$ -mediated inhibition of adenylyl cyclase was not affected. These data suggest that this His.sub.6 fusion protein derived from the carboxyl terminus of BARK1 provides a specific probe for defining  $G(\beta\gamma)$ -mediated processes and for studying the structural features of a  $G(\beta \gamma)$ -binding domain.

L2 ANSWER 10 OF 11 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER: 95190741 EMBASE

DOCUMENT NUMBER: 1995

1995190741

TITLE:

T cell-targeted immunofusion proteins from Escherichia

coli.

AUTHOR:

Better M.; Bernhard S.L.; Williams R.E.; Leigh S.D.; Bauer

R.J.; Kung A.H.C.; Carroll S.F.; Fishwild D.M.

CORPORATE SOURCE:

Xoma Corp., 1545 17th St., Santa Monica, CA 90404, United

States

SOURCE:

Journal of Biological Chemistry, (1995) Vol. 270, No. 25,

pp. 14951-14957.

ISSN: 0021-9258 CODEN: JBCHA3

COUNTRY:

United States
Journal; Article

DOCUMENT TYPE: FILE SEGMENT:

026 Immunology, Serology and Transplantation

029 Clinical Biochemistry

030 Pharmacology

037 Drug Literature Index

LANGUAGE:

English English

SUMMARY LANGUAGE: ENTRY DATE:

Entered STN: 950718

Last Updated on STN: 950718

Fusion proteins between cell-targeting AB domains and cytotoxic proteins should be particularly effective therapeutic reagents. We constructed a family of immunofusion proteins linking humanized Fab, F(ab')2, or single chain antibody forms of the H65 antibody (which recognizes the CD5 antigen on the surface of human T cells) with the plant ribosome-inactivating protein gelonin. We reasoned that such an immunofusion would kill human target cells as efficiently as the previously described chemical conjugates of H65 and gelonin (Better M., Bernhard, S. L., Fishwild, D. M., Nolan, P. A., Bauer, R. J., Kung, A. H. C., and Carroll, S. F. (1994) J. Biol. Chemical 269, 9644-9650) if both the recognition and catalytic domains remained active, and a proper linkage between domains could be found. Immunofusion proteins were produced in Escherichia coli as secreted proteins and were recovered directly from the bacterial culture supernatant in an active form. All of the immunofusion proteins were purified by a common process and were tested for cytotoxicity toward antigen-positive human cells. A 20-60-fold range of cytotoxic activity was seen among the fusion family members, and several fusion

proteins were identified which are approximately as active as effective chemical conjugates. Based on these constructs, immunofusion avidity and potency can be controlled by appropriate selection of antibody domains and ribosome- inactivating protein.

L2 ANSWER 11 OF 11 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights

reserved on STN

ACCESSION NUMBER: 94142823 EMBASE

DOCUMENT NUMBER:

1994142823

TITLE:

. . . .

Functionally active targeting domain of

the  $\beta$ -adrenergic receptor kinase: An inhibitor of  $G(\beta\gamma)$ -mediated stimulation of type II adenylyl

cyclase.

AUTHOR:

Inglese J.; Luttrell L.M.; Iniguez-Lluhi J.A.; Touhara K.;

Koch W.J.; Lefkowitz R.J.

CORPORATE SOURCE:

Department of Medicine, Howard Hughes Medical Institute, Duke University Medical Center, Durham, NC 27710, United

States

SOURCE:

Proceedings of the National Academy of Sciences of the United States of America, (1994) Vol. 91, No. 9, pp.

3637-3641. .

ISSN: 0027-8424 CODEN: PNASA6

COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article

FILE SEGMENT:

029 Clinical Biochemistry

LANGUAGE:

English

SUMMARY LANGUAGE:

English

ENTRY DATE:

Entered STN: 940602

Last Updated on STN: 940602

The  $\beta$ -adrenergic receptor kinase ( $\beta$ ARK) phosphorylates its ABmembrane- associated receptor substrates, such as the  $\beta\text{-adrenergic}$ receptor, triggering events leading to receptor desensitization.  $\beta ARK$ activity is markedly stimulated by the isoprenylated βγ subunit complex of heterotrimeric guanine nucleotide-binding proteins  $(G(\beta\gamma))$ , which translocates the kinase to the plasma membrane and thereby targets it to its receptor substrate. The amino-terminal two-thirds of \$ARK1 composes the receptor recognition and catalytic domains, while the carboxyl third contains the  $G(\beta\gamma)$  binding sequences, the targeting domain. We prepared this domain as a recombinant His6 fusion protein from Escherichia coli and found that it had both independent secondary structure and functional activity. demonstrated the inhibitory properties of this domain against  $G(\beta\gamma)$  activation of type II adenylyl cyclase both in a reconstituted system utilizing Sf9 insect cell membranes and in a permeabilized 293 human embryonic kidney cell system.  $G(i\alpha)$  -mediated inhibition of adenylyl cyclase was not affected. These data suggest that this His6 fusion protein derived from the carboxyl terminus of BARK1 provides a specific probe for defining  $G(\beta\gamma)$  -mediated processes and for studying the structural features of a  $G(\beta\gamma)$ -binding domain.

=> s 12 and trypsin

L4 0 L2 AND TRYPSIN